

(19) World Intellectual Property Organization  
International Bureau



(43) International Publication Date  
16 May 2002 (16.05.2002)

PCT

(10) International Publication Number  
**WO 02/39125 A2**

(51) International Patent Classification<sup>7</sup>: **G01N 35/00**

(21) International Application Number: PCT/US01/29253

(22) International Filing Date:  
18 September 2001 (18.09.2001)

(25) Filing Language: English

(26) Publication Language: English

(30) Priority Data:  
09/677,805 2 October 2000 (02.10.2000) US

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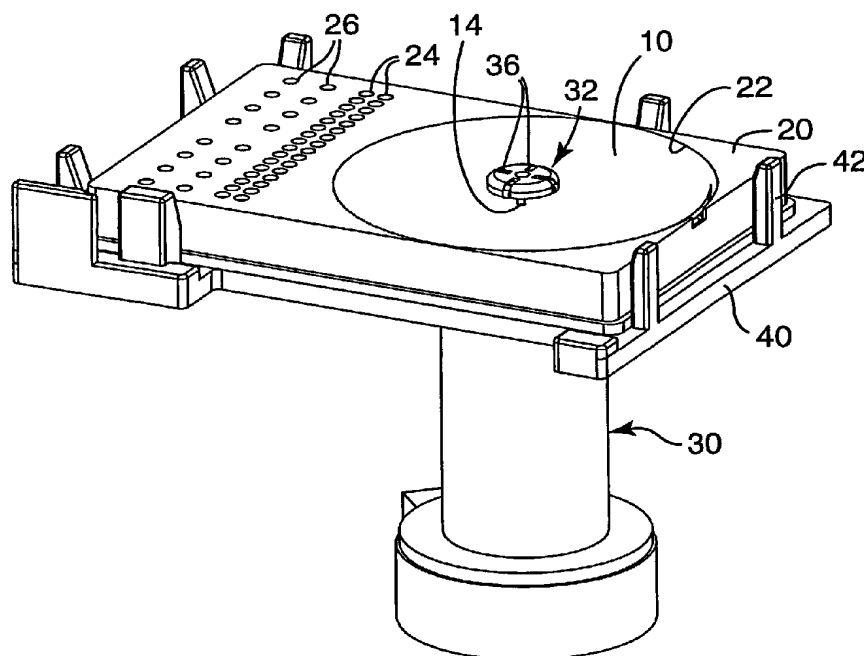
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(81) Designated States (*national*): AE, AG, AL, AM, AT, AT (utility model), AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, CZ (utility model), DE, DE (utility model), DK, DK (utility model), DM, DZ, EC, EE, EE (utility model), ES, FI, FI (utility model), GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SK (utility model), SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW.

(84) Designated States (*regional*): ARIPO patent (GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG).

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(54) Title: SAMPLE PROCESSING APPARATUS, METHODS AND SYSTEMS



(57) Abstract: Apparatus, methods, and systems for processing sample materials that may be presented in a standard microtiter plate are disclosed. The present invention provides a bridge between the standard microtiter plate systems, methods, protocols, etc. with their stationary wells and rotating sample processing devices that allow users to obtain the rapid processing advantages of the newer sample processing devices while retaining the benefits of the standard microtiter plate formats.



WO 02/39125 A2



**Published:**

— without international search report and to be republished  
upon receipt of that report

*For two-letter codes and other abbreviations, refer to the "Guidance Notes on Codes and Abbreviations" appearing at the beginning of each regular issue of the PCT Gazette.*

## SAMPLE PROCESSING APPARATUS, METHODS AND SYSTEMS

### Field of the Invention

5           The present invention relates to sample processing systems and methods. More particularly, the present invention provides apparatus, methods, and systems for processing sample materials.

### Background

10           Many different chemical, biochemical, and other reactions are sensitive to temperature variations. The reactions may be enhanced or inhibited based on the temperatures of the materials involved. Although it may be possible to process samples individually and obtain accurate sample-to-sample results, individual processing can be time-consuming and expensive.

15           Examples of some thermal processes that may be sensitive to temperature variations include, e.g., the manipulation of nucleic acid samples to assist in the deciphering of the genetic code. *See, e.g., T. Maniatis et al. Molecular Cloning, A Laboratory Manual, Cold Spring Harbor Laboratory (1982).* Nucleic acid manipulation techniques include amplification methods such as polymerase chain reaction (PCR); target  
20           polynucleotide amplification methods such as self-sustained sequence replication (3SR) and strand-displacement amplification (SDA); methods based on amplification of a signal attached to the target polynucleotide, such as “branched chain” DNA amplification; methods based on amplification of probe DNA, such as ligase chain reaction (LCR) and QB replicase amplification (QBR); transcription-based methods, such as ligation activated  
25           transcription (LAT) and nucleic acid sequence-based amplification (NASBA); and various other amplification methods, such as repair chain reaction (RCR) and cycling probe reaction (CPR). Other examples of nucleic acid manipulation techniques include, e.g., Sanger sequencing, ligand-binding assays, etc.

                  One approach to reducing the time and cost of thermally processing multiple  
30           samples using such techniques is to use a device including multiple chambers in which different portions of one sample or different samples can be processed simultaneously. Although widely accepted standardized systems have been developed using microtiter

plates having, e.g., 96, 384 or more wells, to speed the processing of multiple sample, even faster sample processing is still desired.

One disadvantage of such devices is, however, their non-standard format as compare to, e.g., the widely accepted standard microtiter plates. As a result, it may be prohibitive in terms of, e.g., equipment costs, test result acceptance, etc. for a facility to abandon the industry standard processes completely and adopt a new test methodology and new equipment.

Another disadvantage of such an approach is in the limited volume available on such devices, which can make cleanup of reaction products difficult due to the limited storage for cleanup solutions.

### **Summary of the Invention**

The present invention provides apparatus, methods, and systems for processing sample materials that may be presented in a standard microtiter plate. More particularly, the present invention provides a bridge between the standard microtiter plate systems, methods, protocols, etc. with their stationary wells and rotating sample processing devices that allow users to obtain the rapid processing advantages of the newer sample processing devices while retaining the benefits of the standard microtiter plate formats.

For example, the rotating sample processing devices can be used to rapidly and accurately perform required processing of the sample materials by virtue of their rotation. That rotation may be used to expose the processing chambers to energy and/or to assist in transporting the sample material to different areas on the processing devices.

In addition, cleanup of the processed sample materials can be performed off of the sample processing device, thereby mitigating any concerns regarding available volume of the sample processing device for any cleaning solutions and materials.

A further advantage of the present invention is its amenability to automation to further enhance speed and/or cost of obtaining the desired results.

In one aspect, the present invention provides an apparatus for processing sample materials, the apparatus including a platform having an upper surface and a lower surface; a plurality of stationary fluid chambers opening at the upper surface of the platform; and retention structure occupying a portion of the upper surface of the platform, wherein the

retention structure is capable of retaining a rotating multi-chambered processing device proximate the upper surface of the platform.

In another aspect, the present invention provides an apparatus for processing sample materials, the apparatus including a platform having an upper surface and a lower surface; a plurality of stationary fluid chambers opening at the upper surface of the platform; retention structure occupying a portion of the upper surface of the platform; and a processing device located within the retention structure proximate the upper surface of the platform, the processing device including a plurality of process chambers, wherein the processing device is capable of being rotated within the retention structure to move the plurality process chambers.

In another aspect, the present invention provides a method of processing sample material by providing a platform having an upper surface and a lower surface, a plurality of stationary fluid chambers opening at the upper surface of the platform, and retention structure occupying a portion of the upper surface of the platform; providing a processing device in the retention structure proximate the upper surface of the platform, the processing device including a plurality of process chambers; providing sample material in a plurality of the plurality of process chambers on the processing device; delivering energy to the process chambers containing sample material to raise the temperature of the sample materials in the process chambers; and rotating the processing device about an axis of rotation within the retention structure while delivering the energy, wherein the temperature of the sample materials in the process chambers is controlled as the processing device rotates to process the sample materials.

In another aspect, the present invention provides a method of processing sample material by providing a platform having an upper surface and a lower surface, a plurality of stationary fluid chambers opening at the upper surface of the platform, and retention structure occupying a portion of the upper surface of the platform, wherein the plurality of stationary fluid chambers are arranged in a rectilinear array on the upper surface of the platform; placing a processing device in the retention structure proximate the upper surface of the platform, the processing device including a plurality of process chambers; positioning at least one of the process chambers on the processing device at a transfer site proximate the upper surface of the platform, wherein the location of the transfer site is

fixed relative to the stationary fluid chambers; loading sample material in a plurality of the plurality of process chambers on the processing device, wherein the process chambers are loaded while positioned at the transfer site; rotating the processing device about an axis of rotation within the retention structure on a spindle extending through a spindle opening  
5 formed through the upper and lower surfaces of the platform; delivering energy to at least some of the plurality of process chambers containing sample material while rotating the processing device to control the temperature of the sample materials in the process chambers, whereby the sample materials are processed; and transferring the sample materials from the process chambers on the processing device to the plurality of stationary  
10 fluid chambers on the platform after processing the sample materials; wherein the sample materials in the process chambers are transferred while the process chambers are located at the transfer site.

In another aspect, the present invention provides a system for processing sample material, the system including a workspace including a processing station; at least one  
15 platform located within the workspace, each platform having an upper surface and a lower surface, a plurality of stationary fluid chambers opening at the upper surface of the platform, and retention structure occupying a portion of the upper surface of the platform; at least one processing device located within the workspace, each processing device including a plurality of process chambers, wherein rotation of the processing device within  
20 the retention structure on the platform moves the plurality process chambers in a circular pattern; a spindle located at the processing station; and a transfer device operative within the workspace, the transfer device capable of transferring sample material from the processing station to another location within the workspace.

These and other features and advantages of the present invention are described with  
25 respect to illustrative embodiments of the invention presented below.

### **Brief Description of the Drawings**

FIGURE 1 is an exploded diagram of one platform and sample processing device according to the present invention.

30 FIGURE 2 is an assembled view of the platform and sample processing device of Figure 1.

FIGURE 3 is an enlarged cross-sectional view of one fluid chamber on the platform including filter material.

FIGURE 4 is a schematic diagram illustrating the positioning of various features of the apparatus on a rectilinear grid.

5       FIGURE 5 is a schematic diagram of a system using the platforms and sample processing devices of the present invention.

FIGURE 6 is an exploded diagram of another platform and sample processing device according to the present invention.

10       FIGURE 7 is an assembled view of the platform and sample processing device of Figure 6.

### **Detailed Description of Illustrative Embodiments of the Invention**

15       The present invention provides apparatus, methods, and systems for processing sample materials that may be presented in a standard microtiter plate. More particularly, the present invention provides a bridge between the standard microtiter plate systems, methods, protocols, etc. and rotating sample processing devices.

20       A portion of one illustrative system is depicted in Figures 1 and 2. The depicted portion of the system includes a sample processing device 10 that may preferably be provided in a disc-shaped format such as that depicted. Examples of suitable disc-shaped devices 10 may be found in, for example, U.S. Provisional Patent Application Serial No. 60/214642 filed on June 28, 2000 and titled SAMPLE PROCESSING DEVICES, SYSTEMS AND METHODS (Attorney Docket No. 55266USA99.003) and U.S. Provisional Patent Application Serial No. 60/237151 filed on even date herewith and titled SAMPLE PROCESSING DEVICES, SYSTEMS AND METHODS (Attorney Docket No. 25 56047USA29). Among the processing devices described in those references, it may be preferable to use those that include features conducive to automated loading and/or unloading as well as automated processing (e.g., tapered inlet ports, seal systems on output chambers, registration systems, etc.). It should further be noted that the devices described in those references may be manufactured according to the principles described in U.S. 30 Provisional Patent Application Serial No. 60/214508 filed on June 28, 2000 and titled

THERMAL PROCESSING DEVICES AND METHODS (Attorney Docket No. 55265USA19.003).

Other processing devices may also be used in connection with the present invention, although they may preferably be modified to incorporate features conducive to automated loading and/or unloading, as well as automated processing. Examples of some potentially useable rotating processing devices include, but are not limited to those described in, e.g., U.S. Patent Nos. 5,160,702 (Kopf-Sill et al.); 5,585,069 (Zanzucchi et al.); 6,030,581 (Virtanen); and 6,063,589 (Kellog et al.); as well as in International Publication No. WO 00/40750 (Orlefors et al.).

A platform 20 is also provided and includes retention structure 22 on its upper surface that is capable of retaining the processing device 10 proximate the upper surface of the platform 20 while the processing device 10 is rotating. The retention structure 22 may preferably be slightly larger than the processing device 10 to allow clearance for the processing device 10 to rotate within the retention structure 22. It may be preferable, but not required, that the retention structure 22 also be in the same general shape as the thermal processing device 10, e.g., circular for disc-shaped processing device 10.

It may be preferred that the retention structure 22 be provided in the form of a cavity as depicted in Figures 1 and 2. On one alternative, the retention structure on a platform 20 may be provided in the form of, e.g., a series of raised posts or protrusions that cooperate to retain the processing device 10 within a desired area on the upper surface of the platform 20.

In one variation, the retention structure may allow for placement and removal of a processing device 10, such as the retention structure 22 illustrated in Figures 1 and 2. As a result, the platform 20 may be used with multiple processing devices 10 that may be located in and removed from the retention structure as needed. In another alternative, such retention structure may allow the user to select a particular processing device 10 for use with the platform 20.

In another variation, the retention structure may be used to retain a captive processing device 10 within a specified area on the upper surface of the platform 20. As used in connection with the present invention, a "captive" processing device means that the processing device cannot be top-loaded into the retention structure without removing a



portion of the platform 20 or somehow distorting the shape of the processing device 10. A captive processing device would be retained within the retention structure even if the platform 20 were oriented such that the upper surface faced the ground.

The depicted platform 20 also includes optional sample fluid chambers 24 that may be used as a staging area for sample materials to be introduced onto the processing device 10. The fluid chambers 24 may also contain one or more reagents that may be used on the processing device 10. The fluid chambers 24 may preferably be arranged in a rectilinear array that is similar in form to arrays of wells found on a standard microtiter plate having, e.g., 96, 384, or more wells.

The platform 20 also includes fluid chambers 26 to be used to receive processed sample material removed from the processing device 10. The fluid chambers 26 may include filter material 27 within the chamber 26 (see Figure 3). The fluid chambers 26 may also include a drain port 28 opening at the lower surface of the platform 20 such that sample material introduced into the chamber 26 may pass through the fluid chamber 26. If filter material 27 is present within the chamber 26, then the sample material is filtered during its passage through the chamber 26.

The fluid chambers 26 on the platform 20 may preferably be arranged in a rectilinear array that is similar in form to arrays of wells found on standard microtiter plates. Further, the fluid chambers 26 are also preferably spaced at intervals compatible with a standard microtiter plate such that wells on a standard microtiter plate located beneath the platform 20 would be aligned with any drain ports of the output wells 26. As a result, processed sample materials removed from the device 10 and transferred to the fluid chambers 26 could be passed through the drain ports of the fluid chambers 26 and into the wells on a standard microtiter plate for additional processing using conventional equipment and procedures.

Also depicted in Figures 1 and 2 is a support carriage 40 including alignment structure 42 for receiving the platform 20. The support carriage 40 may preferably include an opening 44 to allow for passage of processed sample materials from the fluid chambers 26 to a standard microtiter plate located beneath the support carriage 40. The alignment structure 42 may include sloped surfaces as depicted to assist in guiding the platform 20 into position on the support carriage 40.

A spindle 30 is also depicted in Figures 1 and 2 that can be used to rotate the processing device 10 to facilitate processing of the sample materials located thereon. The spindle 30 preferably extends upward through the support carriage 40 and through a spindle opening 23 in the retention structure 22 of the platform 20 (when the plate 20 is located on the carriage 40).

A chuck 32 adapted to mate with the processing device 10 is preferably provided on the end of the spindle 30. The chuck 32 preferably includes a support plate 34 adapted to support the device 10 and at least one, preferably multiple, locking arms 36 in the form of levers that swing outward to retain the device 10 on the chuck 32 as the spindle 30 rotates. The arms 36 may preferably be weighted (or otherwise constructed) such that when the spindle 30 is not rotating, the arms 36 are retracted to allow for placement and removal of the device 10 without hindrance from the arms 36.

In the use of the combination of platform 20 and processing device 10, it may be preferred to position a processing chamber on the processing device 10 at a transfer site proximate the upper surface of the platform 20. A "transfer site" as used herein, is a location at a fixed position relative to the stationary fluid chambers 26. It may be preferred that the stationary fluid chambers 26 and the transfer site 25 be positioned according to a common reference system, e.g., on a rectilinear grid, according to polar coordinates, etc. It may further be preferred that all of the sample fluid chambers 24, stationary fluid chambers 26, transfer site 25 and the location at which loading chambers 16 are positioned for loading be defined by a common reference system.

Referring to Figure 4, one example of a suitable reference system is depicted in which the stationary fluid chambers 26 are located at intersection points on a rectilinear grid, while the transfer site 25 is located at another of the intersection points on the grid. Because the processing device 10 rotates within the retention structure on the platform, different process chambers 12 on the processing device 10 can be indexed into position at the transfer site 25. It may also be preferred that any sample fluid chambers 24, as well as loading chambers 16 on the processing device 10 be positioned at locations defined by a rectilinear array.

Positioning of the processing device 10 to accomplish, e.g., providing the processing chambers 12 on the processing device 10 at the transfer site 25 may be

accomplished by any suitable technique. A variety of registration systems and methods are discussed in, e.g., U.S. Provisional Patent Application Serial No. 60/214,642 filed on June 28, 2000 and titled SAMPLE PROCESSING DEVICES, SYSTEMS AND METHODS (Attorney Docket No. 55266USA99.003).

5           One example of a registration system including complementary structures is depicted in Figures 1 and 2, where one or more of the arms 36 may cooperate with a keyed slot 14 on the device 10 for registering the rotational position of the processing device 10 on the spindle 30 and, therefore, within the retention structure 22 on the platform 20. Such structure can be used to fix the rotational position of the processing device 10 on the  
10 spindle 30, provided for accurate indexing of the processing device 10 to put the various features on the processing device 10 in the desired positions.

The apparatus depicted in Figures 1 and 2 may preferably be used in connection with a variety of transfer devices available from a variety of manufacturers including, but not limited to: Beckman Coulter, Inc. (Fullerton, California), Packard Instrument  
15 Company (Meriden, Connecticut), Tomtec (Hamden, Connecticut), MWG-Biotech Inc. (High Point, North Carolina), and Qiagen (Valencia, California). Such automated transfer devices typically include pick-and place robotic technology and can be outfitted with a processing station including the support carriage 40 and spindle 30 adapted to receive a platform 20 and associated thermal processing device 10.

20           One such system is depicted in Figure 5 and includes a controller 100, transfer device 102, supplies 104 (which may include, e.g., platforms, thermal processing devices, pipette tips, cleanup materials, sample materials, etc.), a processing station 106 (including, e.g., a support carriage 106a, spindle 106b, and source of electromagnetic radiation 106c) and an unloading station 108. All of these components may preferably be located within a  
25 workspace 101.

In use, the processing station 106 can be loaded with the necessary materials from supplies 104 (e.g., a platform, sample materials, processing device, etc.) using the transfer device 102. Sample materials can then be processed at the processing station 106, with the transfer device 102 being used to transfer sample materials to the desired locations (e.g.,  
30 from sample fluid chambers on the platform to loading chambers on a processing device

and/or from process chambers on the processing device to stationary fluid chambers on the platform, etc.).

After processing at the processing station 106, the platform (with processed sample material) may be moved to, e.g., the unloading station 108 using the transfer device 102, where the processed sample materials may be cleaned up using supplies 104 (delivered by the transfer device 102). The sample materials may then be transferred to a standard microtiter plate at the unloading station 108 for further processing of the sample materials.

In some embodiments, it may be preferred to provide the processing station 106 in the form of one of the systems described in, e.g., U.S. Provisional Patent Application Serial No. 60/214,642 filed on June 28, 2000 and titled SAMPLE PROCESSING DEVICES, SYSTEMS AND METHODS (Attorney Docket No. 55266USA99.003) (see, e.g., Figure 9 and the corresponding description in that application) or U.S. Provisional Patent Application Serial No. 60/237151 filed on even date herewith and titled SAMPLE PROCESSING DEVICES, SYSTEMS AND METHODS (Attorney Docket No. 56047USA29).

Furthermore, in another variation the platforms of the present invention may be provided with only a retention structure adapted to receive a rotating processing device, i.e., in the absence of any fluid chambers for staging sample materials or receiving processed sample materials from the processing devices. The rectilinear shape of the preferred platforms allows the rotating processing devices to be used in largely conventional robotic transfer devices. In such a system, it may be desirable, e.g., to provide pipettes or other fluid transfer apparatus that include some filtering structure within them to filter the processed sample materials as they are drawn from the processing devices. The filtered sample materials can then be deposited in the wells of, e.g., standard microtiter plates for conventional processing. Examples of some fluid transfer apparatus that may include materials to clean up samples are described in, e.g., U.S. Patent No. 6,048,457 (Kopaciewicz et al.).

Turning now to Figures 6 and 7, where an alternative apparatus according to the present invention is depicted. The depicted apparatus is useful in those systems and methods in which vacuum is used to remove the processed sample materials from the platform 120. The apparatus includes a platform 120, a vacuum manifold collar 150, and a

manifold base 170 in between which a standard microtiter plate 160 is located when the collar 150 and base 170 are assembled. Together, the vacuum manifold collar 150 and manifold base 170 provide a vacuum manifold useful in extracting sample material from the platform 120 and transferring it to the microtiter plate 160 for processing using conventional processes and equipment.

The platform 120 includes retention structure 122 similar to that described with respect to platform 20 above. A spindle opening 123 is included within the retention structure 122 to receive a spindle adapted to rotate a processing device located therein (not shown).

The platform 120 also includes optional sample fluid chambers 124 that may be used as a staging area for samples to be introduced onto a processing device. The fluid chambers 124 may also contain one or more reagents. The fluid chambers 124 may preferably be arranged in a rectilinear array that is similar in form to arrays of wells on a standard microtiter plate.

The platform 120 also includes stationary fluid chambers 126 to be used to receive processed sample material removed from a processing device. The fluid chambers 126 may include filter material. The fluid chambers 126 on the platform 120 may preferably be arranged in a rectilinear array that is similar in form to arrays of wells found on standard microtiter plates.

Further, the fluid chambers 126 are also preferably spaced at intervals compatible with the spacing of the wells 162 on the microtiter plate 160 such that each of the fluid chambers 126 is located above one of the wells 162 in the microtiter plate 160. It may be preferred that the fluid chambers 126 include extended drain ports 128 adapted to extend at least partially into the microtiter plate wells 162. The extended drain ports 128 may assist in preventing cross-contamination during the transfer of materials from the fluid chambers 126 on the platform 120 to the microtiter plate wells 162.

The vacuum manifold collar 150 preferably includes a seat 152 adapted to receive the lower surface of platform 120. A gasket 156 may preferably be located within the seat 152 to assist in sealing around the lower surface of platform 120 when a vacuum is supplied to the manifold. The collar 150 also includes an opening 154 through which the extended drain ports 128 on the fluid chambers 126 extend when the apparatus is

assembled. The manifold base 170 may also preferably include a gasket 172 to assist in sealing the manifold collar 150 to the manifold base 170.

During use, a vacuum source may be connected to the assembled manifold base 170 and collar 150 to provide a negative pressure that draws processed sample material in the fluid chambers 126 into the microtiter plate wells 162. If the fluid chambers 126 contain filter material, then the processed sample materials in the fluid chambers 126 may be filtered during transfer to the microtiter plate 160. After the transfer is completed, the microtiter plate 160 can be removed and processed using conventional methods and techniques.

One alternative to the use of vacuum in connection with the apparatus depicted in Figures 6 and 7 is the use of positive pressure from above the fluid chambers 126 to force the sample materials in the fluid chambers down through the drain ports 128 and into the microtiter plate 160. In such an apparatus, it may be necessary to provide a pressure manifold that seals about the top surface of the platform 120 around the openings of the fluid chambers 126.

It is to be understood that the above description is intended to be illustrative, and not restrictive. Various modifications and alterations of this invention will become apparent to those skilled in the art from the foregoing description without departing from the scope of this invention, and it should be understood that this invention is not to be unduly limited to the illustrative embodiments set forth herein.

**What is claimed is:**

1. An apparatus for processing sample materials, the apparatus comprising:  
a platform comprising an upper surface and a lower surface;  
a plurality of stationary fluid chambers opening at the upper surface of the  
5 platform;  
retention structure occupying a portion of the upper surface of the platform,  
wherein the retention structure is capable of retaining a rotating multi-chambered  
processing device proximate the upper surface of the platform.
- 10 2. An apparatus according to claim 1, further comprising a spindle opening formed  
through the upper and lower surfaces of the platform, the spindle opening located within  
the retention structure, whereby a spindle is capable of contacting and rotating the multi-  
chambered processing device proximate the upper surface of the platform.
- 15 3. An apparatus according to claim 1, wherein the retention structure comprises a  
cavity formed in the upper surface of the platform.
4. An apparatus according to claim 3, wherein the cavity is circular.
- 20 5. An apparatus according to claim 1, wherein the plurality of stationary fluid  
chambers are arranged in a rectilinear array on the upper surface of the platform.
6. An apparatus according to claim 1, wherein at least some of the plurality of  
stationary fluid chambers further comprise filter material.
- 25 7. An apparatus according to claim 6, wherein the fluid chambers comprising filter  
material further comprise a drain port opening at the lower surface of the platform.
8. An apparatus according to claim 7, wherein each of the drain ports comprises a  
30 drain extension.

9. An apparatus for processing sample materials, the apparatus comprising:  
a platform comprising an upper surface and a lower surface;  
a plurality of stationary fluid chambers opening at the upper surface of the  
platform;

5 retention structure occupying a portion of the upper surface of the platform; and  
a processing device located within the retention structure proximate the upper  
surface of the platform, the processing device comprising a plurality of process chambers,  
wherein the processing device is capable of being rotated within the retention structure to  
move the plurality process chambers.

10 10. An apparatus according to claim 9, further comprising a spindle opening formed  
through the upper and lower surfaces of the platform, the spindle opening located within  
the retention structure, whereby a spindle is capable of contacting and rotating the  
processing device proximate the upper surface of the platform.

15 11. An apparatus according to claim 9, wherein the plurality of stationary fluid  
chambers are arranged in a rectilinear array on the upper surface of the platform.

20 12. An apparatus according to claim 11, wherein at least one of the process chambers  
on the processing device is positioned at a transfer site proximate the upper surface of the  
platform, wherein the location of the transfer site is fixed relative to the stationary fluid  
chambers.

25 13. An apparatus according to claim 12, further comprising complementary registration  
structure on the platform and the processing device, the complementary registration  
structure aligning the at least one process chamber at the location defined by the rectilinear  
array of the stationary fluid chambers when the processing device is stationary.

30 14. An apparatus according to claim 9, wherein the retention structure comprises a  
cavity formed in the upper surface of the platform.



15. An apparatus according to claim 14, wherein the cavity is circular.

16. An apparatus according to claim 9, wherein at least some of the plurality of stationary fluid chambers further comprise filter material.

5

17. An apparatus according to claim 16, wherein the fluid chambers comprising filter material further comprise a drain port opening at the lower surface of the platform.

18. An apparatus according to claim 17, wherein each of the drain ports comprises a drain extension.

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19. An apparatus according to claim 17, further comprising a vacuum manifold.

20. An apparatus according to claim 9, wherein the processing device is captive within the retention structure on the platform.

15

21. A method of processing sample material, the method comprising:

providing a platform comprising an upper surface and a lower surface, a plurality of stationary fluid chambers opening at the upper surface of the platform, and retention structure occupying a portion of the upper surface of the platform;

20

providing a processing device in the retention structure proximate the upper surface of the platform, the processing device comprising a plurality of process chambers;

providing sample material in a plurality of the plurality of process chambers on the processing device;

25

delivering energy to the process chambers containing sample material to raise the temperature of the sample materials in the process chambers; and

rotating the processing device about an axis of rotation within the retention structure while delivering the energy, wherein the temperature of the sample materials in the process chambers is controlled as the processing device rotates to process the sample materials.

30

22. A method according to claim 21, wherein the energy comprises electromagnetic energy.

23. A method according to claim 21, wherein rotating the processing device comprises  
5 extending a spindle through a spindle opening formed through the upper and lower  
surfaces of the platform, the spindle opening located within the retention structure, and  
rotating the processing device using the spindle.

24. A method according to claim 21, further comprising transferring the sample  
10 materials from the process chambers on the processing device to the plurality of stationary  
fluid chambers on the platform after processing the sample materials.

25. A method according to claim 24, wherein the fluid chambers comprise filter  
15 material.

26. A method according to claim 24, further comprising transferring the sample  
material from the fluid chambers of the platform to a microtiter plate comprising a  
plurality of wells.

27. A method according to claim 26, wherein transferring the sample materials  
20 comprises passing the sample materials in the fluid chambers through drain ports opening  
at the lower surface of the platform.

28. A method according to claim 27, wherein the fluid chambers comprise filter  
25 material, and further wherein passing the sample material through the drain ports  
comprises passing the sample materials through the filter material in the fluid chambers.

29. A method according to claim 27, wherein the passing of sample material through  
30 the drain ports is accomplished using vacuum.

30. A method according to claim 29, wherein the vacuum is delivered by placing the lower surface of the platform on a vacuum manifold and drawing a vacuum between the platform and the vacuum manifold.

5 31. A method according to claim 21, wherein the plurality of stationary fluid chambers on the platform are arranged in a rectilinear array on the upper surface of the platform.

32. A method according to claim 31, further comprising positioning at least one of the process chambers on the processing device at a transfer site proximate the upper surface of the platform, wherein the location of the transfer site is fixed relative to the stationary fluid  
10 chambers.

33. A method according to claim 32, further comprising transferring the sample material from the process chamber located at the transfer site to one of the stationary fluid  
15 chambers on the platform after processing the sample materials.

34. A method according to claim 32, wherein the positioning further comprises providing complementary registration structure on the platform and the processing device, the complementary registration structure aligning the at least one process chamber at the  
20 transfer site when the processing device is stationary.

35. A method of processing sample material, the method comprising:  
providing a platform comprising an upper surface and a lower surface, a plurality of stationary fluid chambers opening at the upper surface of the platform, and retention  
25 structure occupying a portion of the upper surface of the platform, wherein the plurality of stationary fluid chambers are arranged in a rectilinear array on the upper surface of the platform;

placing a processing device in the retention structure proximate the upper surface of the platform, the processing device comprising a plurality of process chambers;

positioning at least one of the process chambers on the processing device at a transfer site proximate the upper surface of the platform, wherein the location of the transfer site is fixed relative to the stationary fluid chambers;

loading sample material in a plurality of the plurality of process chambers on the processing device, wherein the process chambers are loaded while positioned at the transfer site;

rotating the processing device about an axis of rotation within the retention structure on a spindle extending through a spindle opening formed through the upper and lower surfaces of the platform;

delivering energy to at least some of the plurality of process chambers containing sample material while rotating the processing device to control the temperature of the sample materials in the process chambers, whereby the sample materials are processed; and

transferring the sample materials from the process chambers on the processing device to the plurality of stationary fluid chambers on the platform after processing the sample materials; where the sample materials in the process chambers are transferred while the process chambers are located at the transfer site.

36. A method according to claim 35, wherein the energy comprises electromagnetic energy.

37. A method according to claim 35, wherein the fluid chambers comprise filter material.

38. A method according to claim 37, further comprising transferring the sample material from the stationary fluid chambers of the platform to a microtiter plate comprising a plurality of wells.

39. A method according to claim 38, wherein the transferring of sample materials from the stationary fluid chambers comprises passing the sample materials in the fluid chambers through drain ports opening at the lower surface of the platform.

40. A method according to claim 39, wherein the fluid chambers comprise filter material, and further wherein passing the sample material through the drain ports comprises passing the sample materials through the filter material in the fluid chambers.

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41. A method according to claim 39, wherein the passing of sample material through the drain ports is accomplished using vacuum.

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42. A method according to claim 41, wherein the vacuum is delivered by placing the lower surface of the platform on a vacuum manifold and drawing a vacuum between the platform and the vacuum manifold.

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43. A method according to claim 35, wherein the positioning further comprises providing complementary registration structure on the platform and the processing device, the complementary registration structure aligning the at least one process chamber at the transfer site when the processing device is stationary.

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44. A system for processing sample material, the system comprising:  
a workspace comprising a processing station;  
at least one platform located within the workspace, each platform comprising an upper surface and a lower surface, a plurality of stationary fluid chambers opening at the upper surface of the platform, and retention structure occupying a portion of the upper surface of the platform;  
at least one processing device located within the workspace, each processing device comprising a plurality of process chambers, wherein rotation of the processing device within the retention structure on the platform moves the plurality process chambers in a circular pattern;  
a spindle located at the processing station; and  
a transfer device operative within the workspace, the transfer device capable of transferring sample material from the processing station to another location within the workspace.

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45. A system according to claim 44, wherein the workspace further comprises an unloading station, and further wherein the transfer device transfers sample material by transferring the at least one platform from the processing station to the unloading station.

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46. A system according to claim 44, wherein the transfer device is capable of transferring sample material from the at least one processing device to the stationary fluid chambers of the at least one platform.

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47. A system according to claim 44, further comprising a plurality of processing devices located within the workspace.

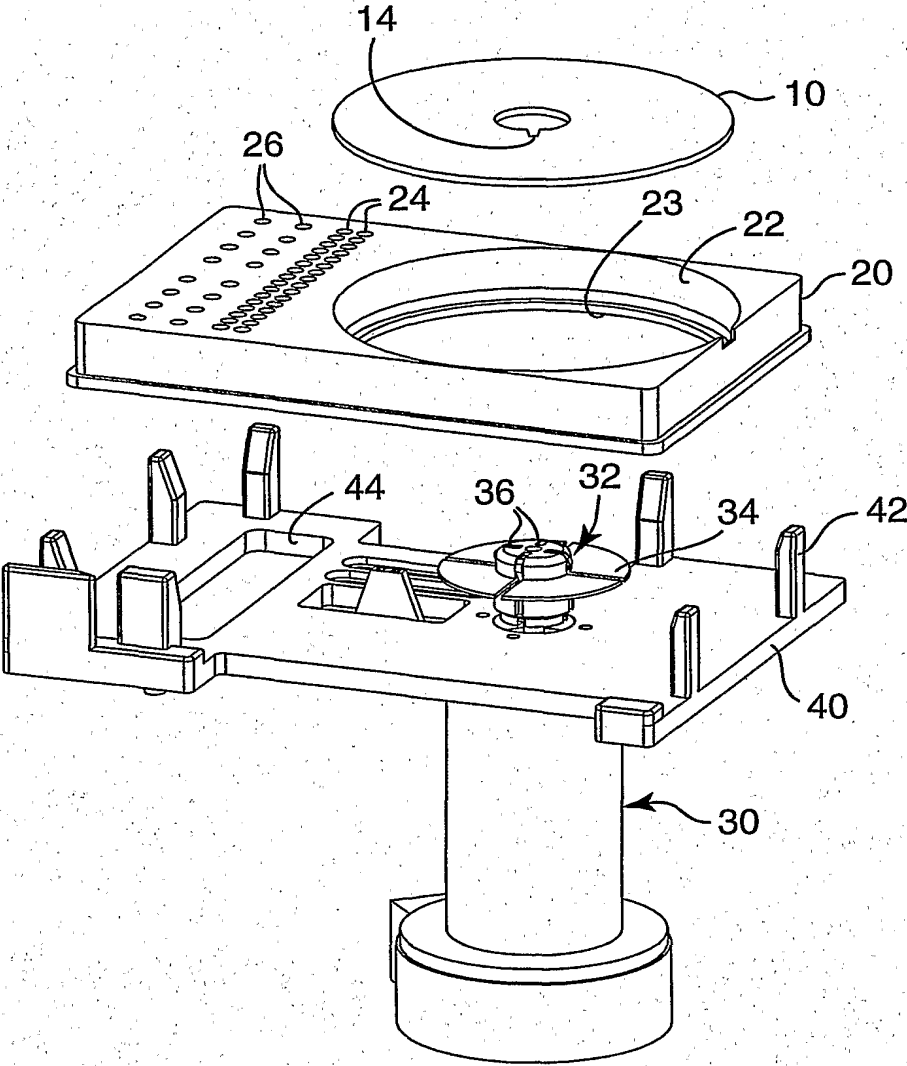
48. A system according to claim 44, further comprising a plurality of platforms located within the workspace.

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49. A system according to claim 44, wherein the at least one platform further comprises a spindle opening formed through the upper and lower surfaces of the platform, the spindle opening located within the retention structure, whereby a spindle is capable of contacting and rotating the processing device proximate the upper surface of the platform when the platform and the processing device are located at the processing station.

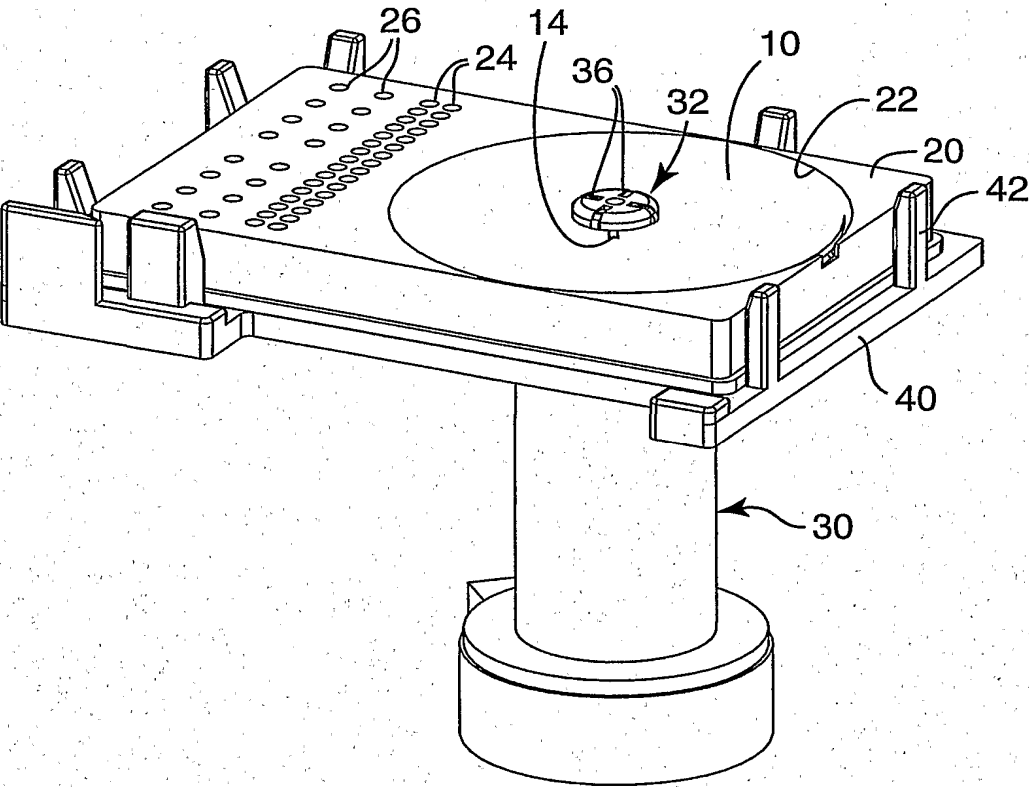
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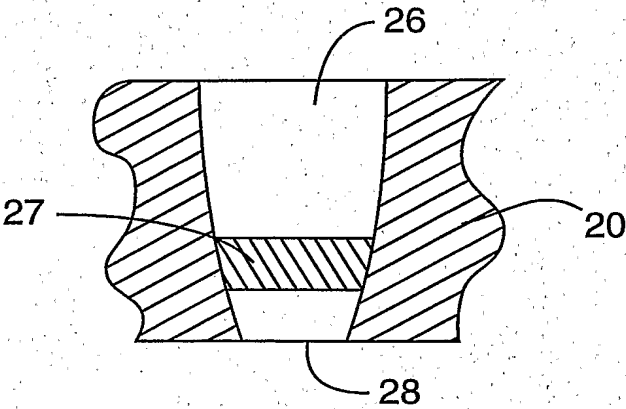


**FIG. 1**

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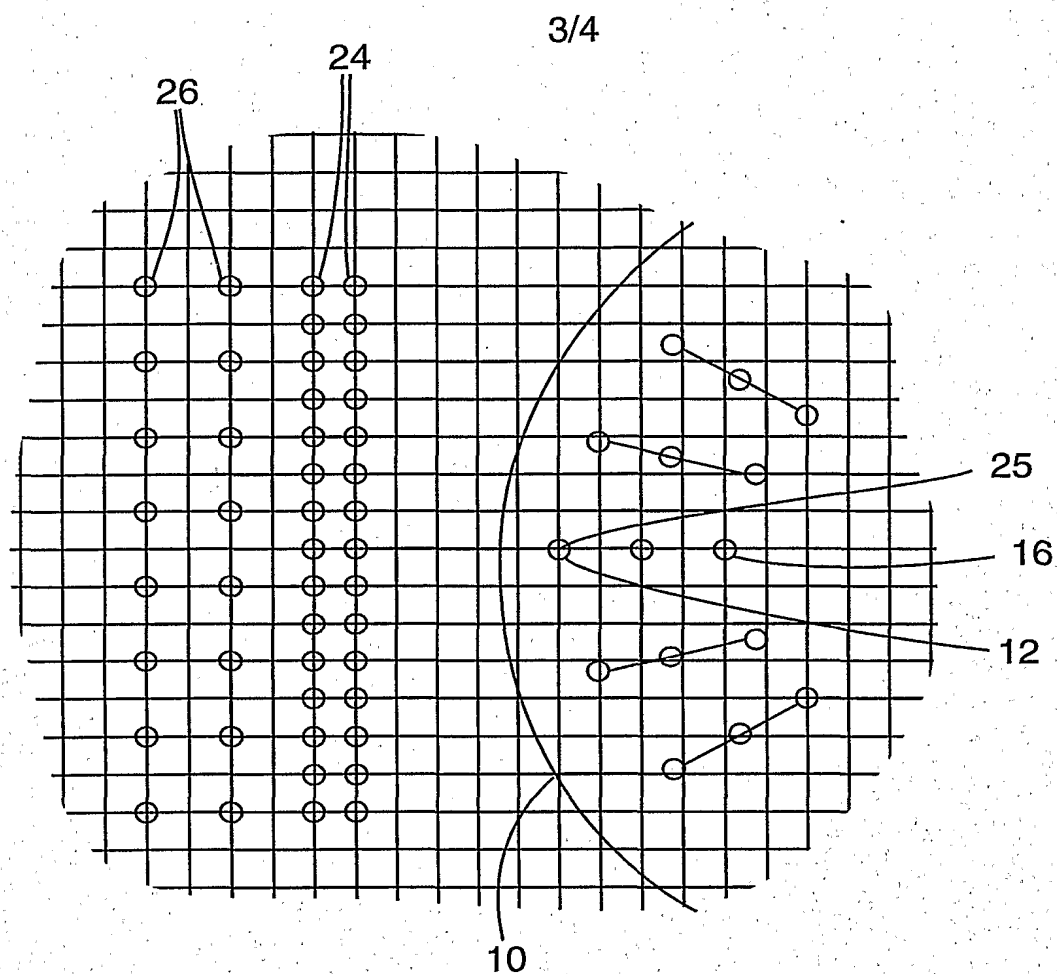
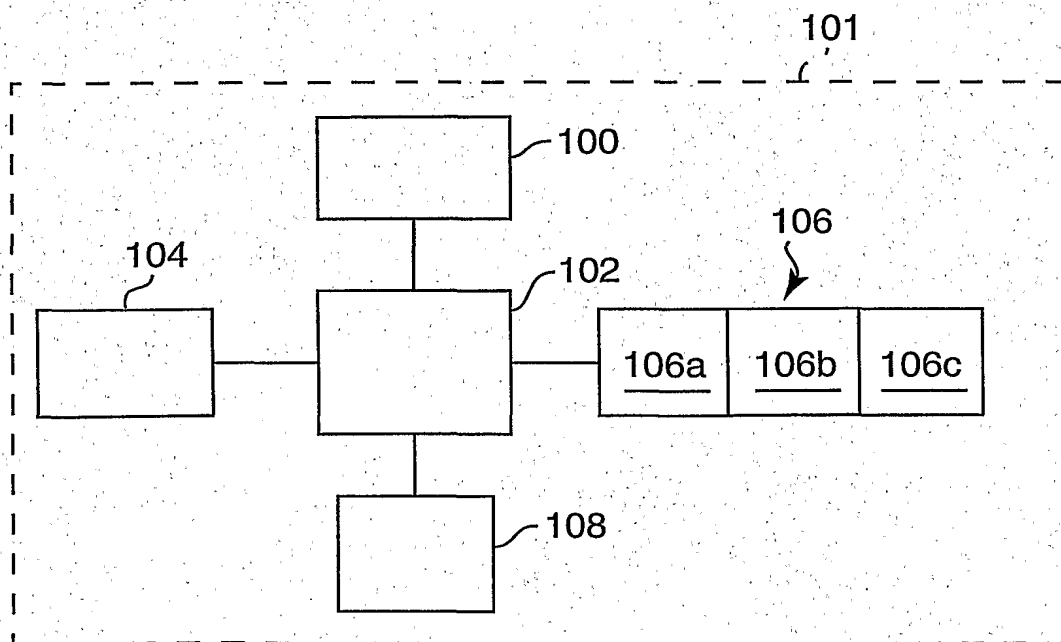


**FIG. 2**

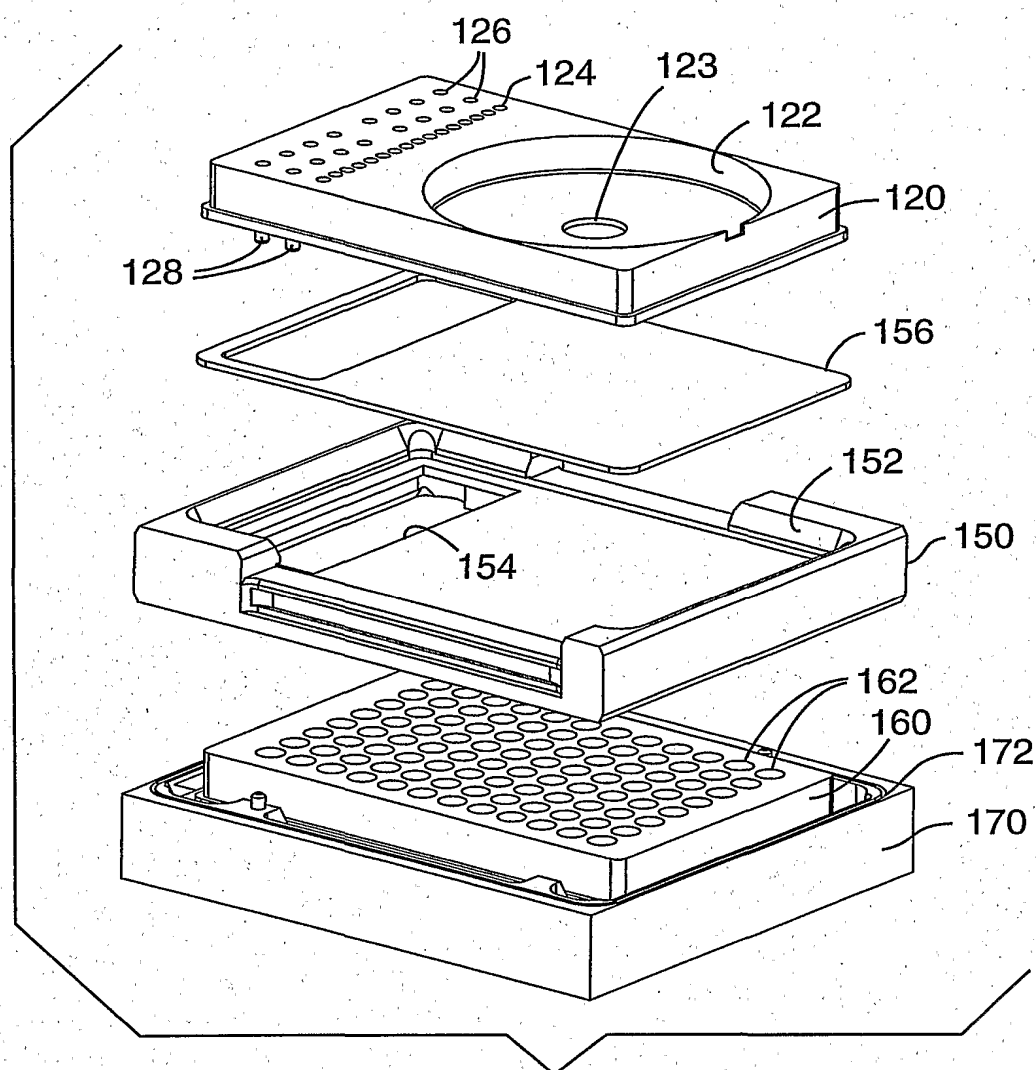
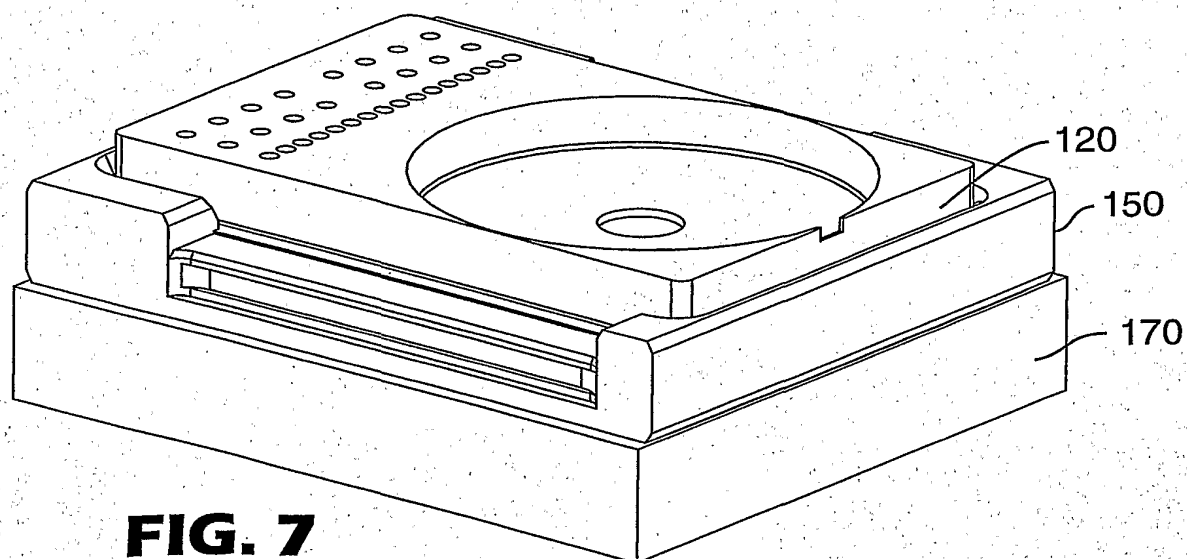


**FIG. 3**



**FIG. 5****FIG. 5**

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**FIG. 6****FIG. 7**

(19) World Intellectual Property Organization  
International Bureau



(43) International Publication Date  
16 May 2002 (16.05.2002)

PCT

(10) International Publication Number  
**WO 02/039125 A3**

(51) International Patent Classification<sup>7</sup>: **G01N 21/07**,  
35/10, B01L 11/00, B04B 5/04

(21) International Application Number: PCT/US01/29253

(22) International Filing Date:  
18 September 2001 (18.09.2001)

(25) Filing Language: English

(26) Publication Language: English

(30) Priority Data:  
09/677,805 2 October 2000 (02.10.2000) US

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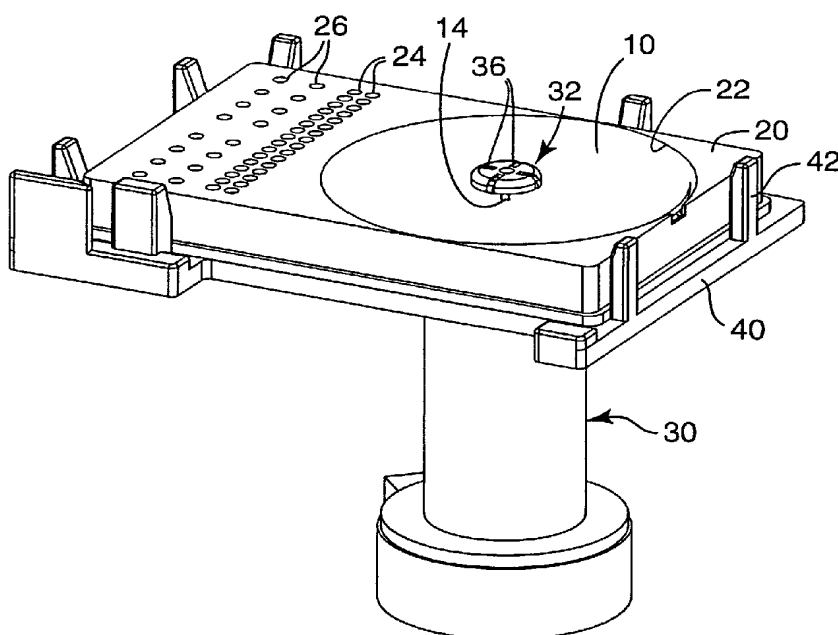
(74) Agents: **GRAM, Christopher, D.** et al.; Office of Intellectual Property Counsel, Post Office Box 33427, Saint Paul, MN 55133-3427 (US).

(81) Designated States (*national*): AE, AG, AL, AM, AT (utility model), AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ (utility model), CZ, DE (utility model), DE, DK (utility model), DK, DM, DZ, EC, EE (utility model), EE, ES, FI (utility model), FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK (utility model), SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW.

(84) Designated States (*regional*): ARIPO patent (GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG).

[Continued on next page]

(54) Title: SAMPLE PROCESSING APPARATUS, METHODS AND SYSTEMS



(57) Abstract: Apparatus, methods, and systems for processing sample materials that may be presented in a standard microtiter plate are disclosed. The present invention provides a bridge between the standard microtiter plate systems, methods, protocols, etc. with their stationary wells and rotating sample processing devices that allow users to obtain the rapid processing advantages of the newer sample processing devices while retaining the benefits of the standard microtiter plate formats.



WO 02/039125 A3



**Published:**

— with international search report

*For two-letter codes and other abbreviations, refer to the "Guidance Notes on Codes and Abbreviations" appearing at the beginning of each regular issue of the PCT Gazette.*

**(88) Date of publication of the international search report:**

27 March 2003

## INTERNATIONAL SEARCH REPORT

International Application No

PCT/US 01/29253

## A. CLASSIFICATION OF SUBJECT MATTER

IPC 7 G01N21/07 G01N35/10 B01L11/00 B04B5/04

According to International Patent Classification (IPC) or to both national classification and IPC

## B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

IPC 7 B04B G01N B01L

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

EPO-Internal, PAJ

## C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
A	WO 00 40750 A (AMERSHAM PHARM BIOTECH AB ;ORLEFORS ANNA EDMAN (SE); ULFENDAHL PER) 13 July 2000 (2000-07-13)  figure 3 page 7, line 7 -page 11, line 19 ---	1-6, 9-16, 20-26, 31-38, 43-49
A	US 4 284 602 A (KELTON ARDEN A ET AL) 18 August 1981 (1981-08-18) figure 1 column 1, line 1 -column 2, line 62 --- -/--	1-49



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Date of the actual completion of the international search

12 December 2002

Date of mailing of the international search report

23/12/2002

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PCT/US 01/29253

## C.(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
A	US 3 902 660 A (BARBER BRUCE WILLIAM) 2 September 1975 (1975-09-02)  the whole document ---	1-6, 9-16, 20-26, 31-38, 43-49
A	PATENT ABSTRACTS OF JAPAN vol. 012, no. 226 (P-722), 28 June 1988 (1988-06-28) & JP 63 019558 A (KONIKA CORP), 27 January 1988 (1988-01-27) abstract ---	1-49
A	US 6 121 054 A (LEBL MICHAL) 19 September 2000 (2000-09-19) figures 6A-6C column 4, line 40 -column 4, line 51 ---	1-37
A	US 5 863 708 A (CHERUKURI SATYAM CHOUDARY ET AL) 26 January 1999 (1999-01-26)  column 1, line 18 -column 2, line 39 column 3, line 63 -column 4, line 32 column 5, line 24 -column 6, line 6 column 7, line 53 -column 8, line 17 column 11, line 8 -column 14, line 35 ---	1-6, 9-16, 20-26, 31-38, 43-49
A	US 6 030 581 A (VIRTANEN JORMA) 29 February 2000 (2000-02-29) the whole document -----	1-49

## INTERNATIONAL SEARCH REPORT

Information on patent family members

International Application No

PCT/US 01/29253

Patent document cited in search report		Publication date	Patent family member(s)	Publication date
WO 0040750	A	13-07-2000	AU 1983800 A WO 0040750 A1 EP 1141401 A1 JP 2002534096 T	24-07-2000 13-07-2000 10-10-2001 15-10-2002
US 4284602	A	18-08-1981	NONE	
US 3902660	A	02-09-1975	GB 1413987 A GB 1423987 A IT 980628 B	12-11-1975 04-02-1976 10-10-1974
JP 63019558	A	27-01-1988	NONE	
US 6121054	A	19-09-2000	AU 751424 B2 AU 1415499 A CA 2309753 A1 EP 1032469 A1 JP 2001523550 T WO 9925470 A1	15-08-2002 07-06-1999 27-05-1999 06-09-2000 27-11-2001 27-05-1999
US 5863708	A	26-01-1999	US 5585069 A AU 705351 B2 AU 4152396 A AU 705659 B2 AU 4233796 A CA 2204912 A1 CA 2205066 A1 EP 0791238 A1 EP 0808456 A1 JP 11500602 T JP 2001509727 T WO 9615450 A1 WO 9615576 A1 US 5681484 A US 5643738 A US 5593838 A US 5846396 A US 5985119 A US 5755942 A US 5858804 A	17-12-1996 20-05-1999 06-06-1996 27-05-1999 06-06-1996 23-05-1996 23-05-1996 27-08-1997 26-11-1997 19-01-1999 24-07-2001 23-05-1996 23-05-1996 28-10-1997 01-07-1997 14-01-1997 08-12-1998 16-11-1999 26-05-1998 12-01-1999
US 6030581	A	29-02-2000	US 2001048895 A1 AU 740195 B2 AU 6346498 A BG 103765 A BR 9808653 A EA 2403 B1 EE 9900377 A EP 0968434 A2 GB 2337113 A ,B HK 1023400 A1 JP 2000515632 T LV 12469 A LV 12469 B NO 994133 A NZ 338017 A SI 20346 A SK 118099 A3	06-12-2001 01-11-2001 18-09-1998 30-06-2000 31-07-2001 25-04-2002 17-04-2000 05-01-2000 10-11-1999 07-12-2001 21-11-2000 20-04-2000 20-09-2000 28-10-1999 30-03-2001 28-02-2001 16-05-2000

# INTERNATIONAL SEARCH REPORT

...formation on patent family members

International Application No

PCT/US 01/29253

Patent document cited in search report	Publication date	Patent family member(s)	Publication date
US 6030581	A	CN 1249816 T	05-04-2000
		HU 0003152 A2	29-01-2001
		LT 99119 A ,B	25-04-2000
		PL 335482 A1	25-04-2000
		TR 9902440 T2	21-02-2000
		WO 9838510 A2	03-09-1998
<hr/>			